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Attorney Docket No. 133114-02CP

Group Art Unit:

3736

Examiner:

Robert L. Nasser

Inventors:

Torok et al.

Serial No.:

09/758,978

Filed:

January 12, 2001

For:

SYSTEM FOR IDENTIFYING PREMATURE RUPTURE OF

MEMBRANE DURING PREGNANCY

Mail Stop Appeal Brief – Patents Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

REPLY BRIEF

Appellants respectfully submit the present Reply to the Examiner's Answer mailed August 1, 2004.

New Points Of Argument - Newly-Cited Reference

In his Answer, the Examiner stated

"[I]f appellant were successful in establishing that bromothymol blue is a known skin irritant, the claims would still be subject to a rejection under 35 U.S.C. 103 as being obvious over Yazaki in view of US Patent 5,660,790 to Lawrence (not of record), as I awrence teaches the equivalence of bromothymol blue and several other chemical pH indicators that change color with the pH levels, so[me] of which are the same as those used by applicant, and accordingly, are non-irritating."
(Examiner's Answer, page 5, lines 5-11)

Appellants respectfully note the following.

PAGE 4/31 * RCVD AT 10/3/2005 2:55:52 PM [Eastern Daylight Time] * SVR:USPTO-EFXRF-6/25 * DNIS:2738300 * CSID:2482581439 * DURATION (mm-ss):07-14

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Application No. 09/758,978 Reply Brief Attorney Docket No. 133114-02CP

A. Bromothymol Blue

In their entered Amendment mailed October 25, 2002, in which all of the independent claims were amended to specify that the pH-sensitive material was "non-irritating to the woman," Appellants stated that in Yazaki the "pH Indication sheet is comprised of a sheet 7 which is wet with bromothymol blue (BTB)...[and]. the sheet 6 is placed underneath the unwoven cloth 5 and, thus, will not be brought to contact to the skin of a wearer." (Page 4, lines 19-23) Conversely, according to Appellants' article, "the pH-sensitive material will be in direct contact with the skin of the woman..." (Page 4, lines 27-28) In their non-entered Amendment mailed June 10, 2003, Appellants stated "Yazaki teaches nothing about a pH-sensitive material that is non-irritating to the wearer." (Page 5, lines 12-13)

Since their first substantive response in the prosecution of the instant application Appellants have directed attention to the fact that their pH-sensitive material is different from that used by Yazaki in that it is non-imitating, and thus was different – by definition – from the effects of bromothymol blue. That this was so must have been understood by the Examiner who never questioned Appellants' position. In any event, Appellants respectfully request that the Board

Earth of the pending independent claims - Claims 2, 7, 13, and 31 - was amended in the Amendment mallest October 25, 2003, Appellants' first substantive amendment, to define the pH material as being non-imitating to the woman. Whether direct contact is specified in the claim or not, each of the independent claims defines the pH-sensitive material as being applied to the article which, by necessity, must be applied to the purface of the article and which, by further necessity, must be worn in direct contact with the wearer. It is not possible for the article to be used in any other way and still achieve its intended purpose of "identifying the premature rupture of a membrane during pregnancy."

Application No. 09/758,978 Reply Brief Atturney Docket No. 133114-02CP

take official notice of the known and readily-ascertainable fact that bromothymol blue is a skin initant.²

characteristics of bromothymol blue may have been submitted in USSN 09/595,594 (co-owned by Appellants herein), now abandoned, which is the parent case to the patent application under appeal. This application which was a continuation of USSN 09/351,875 (now issued as USPN 6,149,590), which was a continuation of USSN 09/120,829 (now issued as USPN 6,126,597). The Appellants are trying to obtain a copy of the file history for USSN 09/595,594 to confirm this one way or the other and will promptly submit a clarifying Supplemental Reply Brief.

B. Lawrence et al.

The Examiner stated that even if Appellants established that bromothymol blue was a known skin irritant, the claims would still be subject to a rejection under 35 U.S.C. Section 103 as being obvious over Yazaki in view of Lawrence et al. "for example, as Lawrence leaches the equivalence of bromothymol blue and several other chemical pH indicators that change color with the pH levels,

² Regarding the taking of "judicial notice." "[a]s to the propriety of the board's laking such notice at all, this court has already previously determined that the Patent Office appellate tribunals, where it is found necessary, may take mittee of facts beyond the record which, while not generally notorious, are capable of such instant and unquestionable demonstration as to defy dispute." In re Ahlert, 424 F.2d 1088, 1091, 165 USPQ 418, 420 (CCPA 1970); see also MPEP Section 2144.03. In this instance, that bromothymol blue may be identified as being a skin initiant may be readily ascertained by, for example, entering "bromothymol blue" and "irritant" into an Internet search engine.

Application No. 09/758,978 Repty Brief Attorney Docket No. 133114-02CP

so[me] of which are the same as those used by applicant, and accordingly, are non-irritating." (Page 5, lines 8-11)

Rather than being applicable as a persuasive Section 103 reference, Appellants respectfully submit that the patent to Lawrence et al. only adds support to the allowability of the application over the prior art. Appellants have argued that Yazakl teaches only the use of an imitating pH-sensitive material, bromothymol blue, a teaching that is contrary to the use of a non-imitating pH-sensitive material as claimed by Appellaes, and is thus inapplicable as a reference for rejection under 35 U.S.C. Section 102(e). With the exceptions only of (1) nitrazine yellow (Appellants' preferred material; please refer to, for example, page 7, lines 14-15, of the application as filled) and (2) bromochlorophonol blue³, the toxicological properties of which "have not been fully investigated," each of the indicators list in Lawrence et al. (col. 5, lines 50-67) can be irritating to the skin:

Indicator	CAS Number	Potential Health Effect	Exhibit No.
Acid blue 92 Acid blue 29	3861-73-2 5850-35 1	"Skin: May cause skin irritation." "Skin: May be harmful if absorbed through the skin. May cause irritation."	2
Acid alizarin violet N Bromophenol	2092-55-9 155-39-9	"Irritant." "Personal protection – Safety glasses, gloves." "Primary irritant effect: on the skin: May cause irritation."	└
Bromochloro- phenol blue	102185-52-4	"Skin: No Information regarding skin irritation and other potential effects	5

³ Even if non-irritating, the pH range of this indicator is between 3.0 arm 4.0, making it unsuitable for use in detecting armiotic fluid which has a pH range of 7.0-7.5. (Please see attached Exhibit A).

Application No. 09/758,978

Reply Brief

Attorney Docket No. 133114-02CP

		was tound	
Bromocresul	76-60-8	"Irritating to eyes and skin."	
green {	4400 00 0	*Skin. Causea skin irritation.*	<i>'</i>
Chlorophenol red Bromocresol purple	4430-20-0 115-40-2	Primary irritant effect on the skin Irritant to skin and mucous	
Alizarin complexone	3952-7A-1	*Skin: May cause skin imitation. *	9
dehydrate Alizarin red S monohydrate	130-22-3	"Irritating to eyes, respiratory system and skin." "Causes skin irritation." "May be harmful if absorbed through the skin."	10
Bromothymol	76-59-6	"Tuxicology: May act as irritant."	11
blue Brilliant yellow	3051-11-4	"Contact with eyes and skin may result in initiation."	12
Phenoi red	34487-61-1	"Skin Contact: May cause initation	13
Cresol red	1733 12-6	"Primary irritant effect: on the skin:	
m-cresol purple	2303-01-7	"Skin Contact: May cause imitation	16 _
Thymol blue	76-81- <u>9</u>	"Skin: May cause irritation."	T'10 ——

Appellants respectfully submit that the patent to Lawrence et al., if combined with Yazaki, would only serve to further support Appellants' position that the relevant prior art teaches only the use of irritating indicators and teaches away from the use of non-Initaling indicators.

Because Yazaki does not disclose what is claimed, because Yazaki would not otherwise render the present invention obvious, and because the combination of Yazaki and Lawrence et al. reinforces the position that the prior art fails to teach, suggest, or otherwise render obvious the use of a non-irritating pH-sensitive material for use in "identifying the premature rupture of a membrane Application No. 09/758,978 Reply Brief Altorney Docket No. 133114-02CP

during pregnancy," Appellants respectfully submit that the Examiner's rejection on the basis of prior art is in error and should be reversed.

CONCLUSION

Appellants respectfully submit that the appealed application is in condition for allowance and that the Examiner's outstanding rejections of this application are in error and should be reversed.

Appellants do not believe that an Oral Flearing will be necessary unless deemed so by the Board.

Respectfully submitted,

Thomas T. Moga Registration No. 34,881 Attorney for Appellants

BUTZEL LONG 100 Bloomfield Hills Parkway Suitc 200 Bloomfield Hills, MI 48304 (248) 258-4496

Dated: October 3, 2005

ConstructionWork Construction Work Center

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pH Indicators

Indicator	Aeid Color	Low pH	High pH	Base Color
Methyl vlolet	velluw	0.0	1.5	plue
мещун чолог . Cresol red (1st галув)	red	0.0	1.8	yellow
	vellow	0.0	1.6	pino
· Orystal violet	yellow	0.2	1,8	pin o Guseru
Malachite grash	yellow	0.2	1.8	bluc
Mothyl green	red	1.2	2.4	yellow
_ Matanli yelluw	red	1.2	2.8	yollow
m-Crosol purple (1st range)	red	1.2	2.8	yellow
Metacresol purple (1st range)	rod	12	2.8	yellow
Thymol blue (1st range)	orance .	1.4	2.8	yellow
4-o-Tulylazo-o-toluidine	teg realite	1.4	3.2	yallow
Orange IV (Tropentin 00)	caluriess	1.7	4.4	yellaw
2,6-Dintrophenol	red	1.9	3.3	yellow
Велгуі пғанұв		2.0	4.7	vellow
2,4-Dinitrophenol	coloriesa violat	2.2	4.2	rect
- Benzopurputine 48		2.2 2.9	4.0	vellow
p-Dimothylaminoszobenzens	red 	3.0	4.B	numie
Bromochlæophendi bluc	yellow		4.6	bhic
Bromophenot blue	yellow	3.0	5.0	red
Congo rod	blue	3.0	4,4	yellow
Methyl crange	red	3.1	4.6	yellow
Ethyl grange	red	3.4	5.4	blue
Bromncresol green	yellow	3.8		viulal
Rosazurin	orange	3.8	ñ.4	
2,5-Dinotrophenol	coloriess	4.0	5.8	yellow
Mathyl red	।ध्य	4.2	6.3	Arrigna
Azollinin (litmus)	red	4.4	6.6	euld
Agraniu ted 2	yellow	4.6	6.0	red
Propyl rod	red	4.6	8.6	Asilom
Chlorophenol red	yallow	4.6	7.0	red
p-Nitrophenol	coluntess	4.7	7.9	yellow
Bromophenol red	yellow	4.8	8.8	purple
Bromocrosol purple	yellow	5.2	6.8	purple
p-Nitrophenol	coloriess	5.4	6.6	yellow
Bromothymni blue	yellow	6.0	7.8	blue
Brījilant yellow	yellow	6.6	7.8	ren

Exhibit - A USSN 09/758,978 Torok et e/.

New Page 2

This is G o o g t e's cache of http://www.selsc.noss.gov/HTMLdocs/coomassic blue.htm. G o o g I e's cache is the snapshot that we took of the page as we crawled the web. The page may have changed since that time, Click here for the current page without highlighting. To link to or bookmark this page, use the following url: nccp://www.google.com/search? Q=DBclis:cBBhbVBsrokJ:www.8sfsc.pose.gov/wrMlcncs/copmousic_blue.hem+3s61-73-2+8kinehl-sn

Chogle is not officiated with the ambors of this page nor responsible for its content.

These search terms have been highlighted: 3861 73 2 skin

```
**** EECTION 1 - CHEMICAL PRODUCT AND COMPANY IDENTIFICATION ****
```

MEDS Name: Acid Blue 92

Catalog Numbers:

AC198630000, AC198630050, AC198631000

Synonymo:

2,7-Naphthalenedisulfonic

acid,4-((4-anilino-5-sulto-)-naphthyl)-5 hydroxy-trisudium salt ;

C.I. 13390 ; Coomassis Blue

Company Identification (Europe): Acros Organids EVBA

Janssen Pharmaceuticalaan 3a

2440 Geel, Belgium

Company Identification (USA):

Acros Organica One Reagent Lane

Pairlawn, N.T 07410 800-ACROS-03

For information in North America, call;

0032(0) 14575311

For information in Enrope, call: For emergencies in the US, call CHEMTREC: 800-424-9300

For emergencies in Europe, call:

0032(0) 14575299

**** SECTION 2 - COMPOSITION, INFORMATION ON INGREDIENTS ****

		··	F
	Chemical Name		
CAS#		100.0	223-378-0
	Acid Blue 92		

Hazard Symbols: None Misted. Risk Phrases: None Listed.

**** SECTION 3 - WAZARDS IDENTIFICATION ****

EMBRGENCY OVERVLKW

Appearance: Dark Blue Powder. Caution! The toxicological properties of this material have not been fully investigated. May cause respiratory and discative tract irritation. May cause eye and skin irritation. Target Organs: None.

Potential Health Effects

May cause eye irritation.

May cause skin irritation.

Ingestion:

Exhibit - 1 ussn 09,758,978 Torok et et.

Page 2 of 9

MATERIAL CAPETY DATA SHEET

HELENA LABORATORIES 1530 Lindbergh Dr./P.O. Box 752 Beaumont, TX 77704-0752 USA Toll Free 800-231-5663

DATE PREPARED: 1/17/1992 REVISION:

1. IDENTIFICATION:

NAME: ACID BLUE 29

CAS #: 5850-35-1

MOLECULAR FORMULA: C22H16N60952

2. HAZARDOUS INGREDIENTS:

EXPOSURE LIMITS CHEMICAL NAME/COMMON NAME OSHA PET. ACCIH TLV

CAS #

NA 100.0 ACID BLUE 29

5050-35-1

3. PHYSICAL DATA:

SPECIFIC GRAVITY (H2O = 1): NA BOILTING POINT: NA

MELTING POINT: NIF VAPOR PRESSURE (MM HG.): NA

SOLUBILITY IN WATER: YES VAPOR DENSITY (AIR - 1): MA

APPEARANCE AND ODOR: BLACK POWDER

MA = Not Applicable

NE - Not Established NIP = No Information Found

NL = Not Listed

Page 3 of 9

MSDS: ACID BLUE 29

4. HEALTH HAZARD DATA:

PRIMARY ROUTE (S) OF ENTRY:

INHALATION? YES

SKIN? YES INGESTION?

YES

CARCINOGENICITY: NTP? NO

IVECS NO

ACUTE EFFECTS OF OVEREXPOSURE:

RYES:

MAY CAUSE IRRITATION.

SKIN:

MAY BE HARMFUL IF ABSORBED THROUGH THE SKIN. MAY

CAUSE IRRITATION.

INHALATION: MAY BE HARMFUL IF INHALED. MAY CAUSE IRRITATION.

INGESTION: MAY BE HARMFUL IF SWALLOWED. MAY CAUSE IRRITATION.

CHRONIC EFFECTS OF EXPOSURE: THE TOXICOLOGICAL PROPERTIES HAVE NOT

BEEN THOROUGHLY INVESTIGATED.

TOXICITY DATA: NO DATA AVAILABLE.

MEDICAL CONDITIONS AGGRAVATED BY EXPOSURE: MIF

EMERGENCY AND FIRST AID PROCEDURES:

EYE CONTACT: IN CASE OF EYE CONTACT, FLUSH WITH COPIOUS AMOUNTS OF WATER FOR AT LEAST 15 MINUTES. ASSURE ADEQUATE FLUSHING BY SEPARATING THE EYELIDS WITH FINGERS.

CALL A PHYSICIAN.

SKIN CONTACT: IN CASE OF SKIN CONTACT, FLUSH WITH COPIOUS AMOUNTS OF WATER FOR AT LEAST 15 MINUTES. REMOVE

CONTAMINATED CLOTHING AND SHORE. CALL A PHYSICIAN.

INHALATION:

IF IMHALED, REMOVE TO FRESH AIR. IF BREATHING

BECOMES DIFFICULT, CALL A PHYSICIAN.

ingration:

IF SWALLOWED, WASH OUT MOUTH WITH WATER PROVIDED

PERSON IS CONSCIOUS. CALL A PHYSICIAN.

5. FIRE AND EXPLOSION HAZARD:

FLASHPOINT (METHOD USE): NA

PLAMMABLE LIMITS:

NA LEL

UEL MA

EXTINCUISHING MEDIA: WATER SPRAY, CARBON DIOXIDE, DRY CHRMICAL POWDER OR APPROPRIATE FOAM.

******************************* 5. PIRE AND EXPLOSION HAZARD (CONT):

NA

PAGE 13/31 * RCVD AT 10/3/2005 2:55:52 PM [Eastern Daylight Time] * SVR:USPTO-EFXRF-6/25 * DNIS:2738300 * CSID:2482581439 * DURATION (mm-ss):07-14

Safety (MSDS) data for acid alizarin violet N



General

Synonyms: 4-hydroxy-3-((2-hydroxy-1-naphthalcnyl)azo)-benzenesulfonic acid monosodium salt, acid alizarin violet B, acid alizarin violet, acid chrome violet K, acid chrome violet N, chromal violet B, C.I. 15670, yodochrome violet B, C.I. mordant violet 5, numerous further names

Use: indicator, stain

Molecular formula: C₁₆li₁₂N₂O₅S

CAS No: 2092-55-9 FINECS No: 218-246-4

Physical data

Appearance: red powder

Melting point:

Boiling point:

Vapour density:

Vapour pressure:

Density (g cm⁻³):

Flash point:

Explosion limits:

Autoignition temperature:

Water solubility:

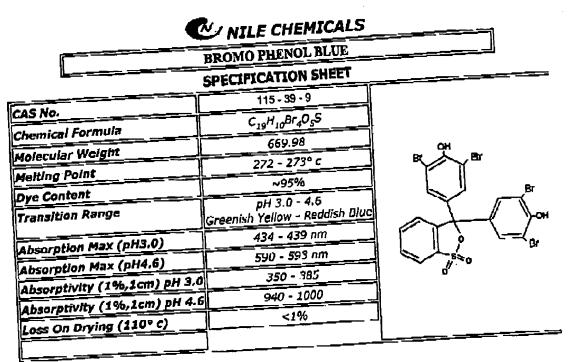
Stability

Stable. Incompatible with strong exidizing agents.

Toxicology

lrritant.

Exhibit - 3 Ussa 09/758,978 Torok et ef. Bronto phonol Blue



PRODUCT LIST HOME

BRIEF MSDS

Hazards identification

Hazard description: Not applicable
 Information pertaining to particular dangers for man and environment: - Not applicable

Tuxicological information

Acute texicity:
 Primary irritant effect:
 on the skin: May cause irritation
 on the eye: May cause irritation
 sensitization: No sensitizing effects known.

Transport information

Not a hazardous material for transportation,

• DOT regulations:

Hazard class:

Land transport ADR/RID (cross-border)

ADR/RID class:

Maritime transport TADG:

Exhibit - 4 ussn 08/758,978 Torok ef ef.

Material Safety Data Sheet Bromochlorophenol blue, water soluble

ACC# 42758

Section 1 - Chemical Product and Company Identification

MSDS Name: Bromochiorophenoi blue, water soluble

Catalog Numbers: AC199680000, AC199680010, AC199680050

Synonyms: None

Company Identification:

Acros Organics N.V. One Reagent Lane Fair Lawn, NJ 07410

For information in North America, call: 800-ACROS-01 For emergencies in the US, call CHEMTREC: 800-424-9300

Section 2 - Composition, Information on Ingredients

ı		Percent	EINECS/ELINCS
1	CAS# Chemical Name	100.0	219-861-0
	2553-71-1 Bromochlorophenol blue	20012	

Hazard Symbols: None listed. Risk Phrases: None listed.

Section 3 - Hazards Identification

EMERGENCY OVERVIEW

Appearance: black powder. Caution! The toxicological properties of this material have not been fully investigated. Target Organs: None.

Eye: No information regarding eye irritation and other potential effects was found. Skin: No information regarding skin irritation and other potential effects was found. Ingestion: The toxicological properties of this substance have not been fully investigated. Inhalation: The toxicological properties of this substance have not been fully investigated. Chronic: No information found.

Section 4 - First Aid Measures

Eyes: Flush eyes with plenty of water for at least 15 minutes, occasionally lifting the upper and lower eyellds. Get medical aid Immediately.

Skin: Flush skin with plenty of water for at least 15 minutes while removing contaminated clothing

and shoes. Get medical aid it irritation develops or persists.

Ingestion: If victim is conscious and alcrt, give 2-4 cupfuls of milk or water. Get medical aid

JSSN 09/758,976 Torok et al. Exhibit

Bromocresol Green

MATERIAL SAFETY DATA SHEET

SECTION 1. ------- CHEMICAL IDENTIFICATION------

CATALOG #:

02559

BROMOCRESOL GREEN SOLUTION, READY-TO-USE

SECTION 2. ---- COMPOSITION/INFORMATION ON INGREDIENTS -----

CAS#: 76-60-8

EC NO: 200-972-8

HAZARDOUS INGREDIENTS

CONTAINS 2-PROPANOL (ISOPROPYL ALCOHOL), CHEMICAL ABSTRACTS REGISTRY

NUMBER 67-63-0.

LAREL PRECAUTIONARY STATEMENTS

FLAMMABLE (USA)

HIGHLY FLAMMABLE (EU)

IRRITANT

IRRITATING TO EYES AND SKIN.

VAPORS MAY CAUSE DROWSINESS AND DIZZINESS.

TARGET ORGAN(S):

NERVES

KIDNEYS

SKIN IRRITANT.

KEEP CONTAINER TIGHTLY CLOSED.

KEFP AWAY FROM SOURCES OF IGNITION - NO SMOKING. '

AVOID CONTACT WITH SKIN AND FYES.

IN CASE OF CONTACT WITH EYES, RINSE IMMEDIATELY WITH PLENTY OF

WATER AND SEEK MEDICAL ADVICE.

SECTION 4. ------FIRST-AID MEASURES------

IF SWALLOWED, WASH OUT MOUTH WITH WATER PROVIDED PERSON IS CONSCIOUS.

CALL A PHYSICIAN.

IF INHALED, REMOVE TO FRESH AIR. IF NOT BREATHING GIVE ARTIFICIAL

RESPIRATION. IF BREATHING IS DIFFICULT, GIVE OXYGEN.

IN CASE OF CONTACT, IMMEDIATELY WASH SKIN WITH SOAP AND COPIOUS

AMOUNTS OF WATER.

IN CASE OF CONTACT, IMMEDIATELY FLUSH EYES WITH COPIOUS AMOUNTS OF

Extribit - 6 USSN 09/758,978 Torok et al.

Material Safety Data Sheet Chlorophenol Red, Indicator Grade

ACC# 72272

Section 1 - Chemical Product and Company Identification

Catalog Numbers: AC190040000, AC190040050, AC190040100, AC9532978, AC9655454,

AC9681524, XXAC19004-40, XXAC1900410K, XXAC1900415K

Synonyms: CPR; 3'-3'-Dichlorophenylsulfonaphthaleln

Company Identification:

Fisher Scientific 1 Reagent Lane

Fair Lawn, NJ 07410

For Information, call: 201-796-7100 Emergency Number: 201-796-7100

For CHEMTREC assistance, call: 800-424-9300

For International CHEMTREC assistance, call: 703-527-3887

Section 2 - Composition, Information on Ingredients

11		
iι		EINECS/ELINCS
-	Percent	ETHECOVERY
	Name	224 619-2
	CAS# Chemical Wallie ca 100	224 013-2
-1		
- 1-	4430-20-0 Chlorophenol Red	
- 1	4430-20-0 Chlorophenol Ked	

Hazard Symbols: XI Risk Phrases: 36/38

Section 3 - Hazards Identification

EMERGENCY OVERVIEW

Appearance: brown solid. May cause respiratory and digestive tract irritation. The toxicological properties of this material have not been fully investigated. Causes eye and skin irritation.

Warningi

Target Organs: No data found.

Potential Health Effects

Eye: Causes cye Irritation.

Ingestion: May cause gastmintestinal irritation with nausea, vomiting and diarrhea. The

toxicological properties of this substance have not been fully investigated.

Inhalation: May cause respiratory tract irritation. The toxicological properties of this substance

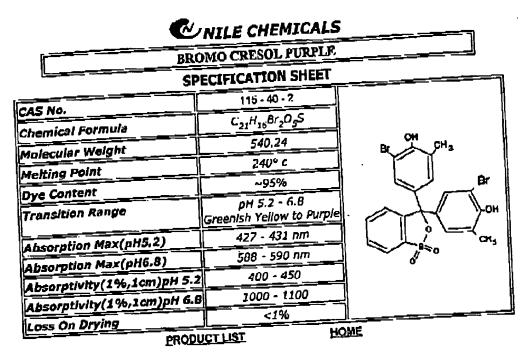
have not been fully investigated.

Chronic: No Information found.

Section 4 - First Aid Measures

Exhibit - 7 USSN 09/758,978 Torok et al.

Bromo cresol purple



Hazards identification

• Hazard description: Xi Irritant.

Information pertaining to particular dangers for man and environment
R 36/37/38 Irritating to eyes, respiratory system and skin.

Toxicological information

• Acute toxicity:

Primary irritant effect:

on the skin: Irritant to skin and mucous membranes.

on the sye: Irritating effect.

Sensitization: No sensitizing effects known.

Transport information

Not a hazardous material for transportation.

• DOT regulations:

Hazard class:

None

Land transport ADR/RID (cross-border)
ADR/RID class:

- . Maritime transport IMDG:
- . IMDG Class:

Notic

Exhibit - 8 ussn 09/758,978 Torok et al.

Material Safety Data Sheet Alizarin Complexone Dihydrate, Indicator Grade

ACC# 96598

Section 1 - Chemical Product and Company Identification

MSDS Name: Alizarin Complexone Dihydrate, Indicator Grade

Catalog Numbers: AC155R30000, AC155830010, AC155830050

Synonyms: (3,4-Dihydroxy-2-Anthraquinonyl)methylimlnodiacetic Acid Dihydrate; Alizarin

Complexon; Alizarine Fluorine Blue.

Company Identification:

Acros Organics N.V. One Reagent Lane Fair Lawn, NJ 07410

For information in North America, call: 800-ACROS-01 For emergencies in the US, call CHEMTREC: 800-424-9300

Section 2 - Composition, Information on Ingredients

		INECS/ELINCS
Chemical Name	Percent	
	>99,00	223-544-2
3952-78-1 Alizarin Complexone Dihydrate		

Hazard Symbols: None listed. Risk Phrases: None listed.

Section 3 - Hazards Identification

EMERGENCY OVERVIEW

Appearance: orange-yellow powder. Caution! The toxicological properties of this material have not been fully investigated. May cause eye and skin irritation. May cause respiratory and digestive tract irritation.

Target Organs: No data found.

Eye: Dust may cause mechanical Irritation. The toxicological properties of this material have not

Skin: May cause skin irritation. The toxicological properties of this material have not been fully

Ingestion: May cause digestive tract disturbances. The toxicological properties of this subs

have not been fully investigated.

Inhalation: Dust is irritating to the respiratory tract. May cause respiratory tract irritation.

toxicological properties of this substance have not been fully investigated.

Chronic: No information found.

Section 4 - First Aid Measures

USSN 09/756,978 Exhibit Torok Alizarin Red S, monohydrate, certified

Valid 11/2001 - 01/2002

Aldrich Chemical Co., Inc. 1001 West St. Paul Milwaukee, WI 53233 UGA Tel: 414-273-3950

:.;

MATERIAL SAFETY DATA SHEET

```
SECTION 1. - - - - - - CHEMICAL IDENTIFICATION- - - - - -
                        119963
                        ALIZARIN RED S MONOHYDRATE, CERTIFIED
    CATALOG #:
               - - COMPOSITION/INFORMATION ON INGREDIENTS - - - -
    name:
SECTION 2. - -
    CAS #:NONE
    MY: C14H807$
    EC NO: 204-981-8
   ACID RED ALIZARINE * AHCUCUINONE RHO E * ALIZARIN CARMINE (BIOLOGICAL
  SYNONYMS
    STAIN) * ALIZARINE CARMINE INDICATOR * ALIZARINE RED A * ALIZARINE
  PED AS - ALIZARINE RED INDICATOR - ALIXARINE RED S (BIOLOGICAL STAIN)
    * ALIZARINE RED S SODIUM SALT * ALIZARINE RED SW * ALIZARINE RED SZ *
    ALIZANINE RED W * ALIZARINE RED WA * ALIZARINE RED FOR WOOL *
    ALIZARINE RED WS . ALIZARINE S . ALIZARINE S EXTRA CONC. A EXPORT .
    ALIZARINE S EXTRA PURE A * ALIZARIN RED S * ALIZARINROT-S (GERMAN) *
    ALIZARIN S * ALIZARINGULFONATE * CALCOCUROME ALIXARINE RED SC *
    CARNELTO RUBINE LAKE * CHROME RED ALIZARINE * C.I. 58005 * C.I.
    MORDANT RED 3 * DIAMOND RED W * 9,10-DINYDRO-3,4-DINYDROXY-9,10-DIOXO-
     2-ANTHRACENERULFONIC ACID MONOCODIUM SAIT * EXT D AND C RED NO. 7 *
  FENAKROM REI) W * MITSUI ALIVARINE RED S * OXANAL FAST RED SW * SODIUM
     ALIZARINESULFONATE = SODIUM ALIZARIN-3-SULFONATE *
 SECTION 3. - - - HAZARDS IDENTIFICATION - -
   LAMEL PRECAUTIONARY STATEMENTS
     IRRITANT
     IRRITATING TO EYES, RESPIRATORY SYSTEM AND SKIN.
     CAUSES SEVERE IRRITATION.
     IN CASE OF CONTACT WITH BYES, RINSE IMMEDIATELY WITH PLENTY OF
     WATER AND SEEK MEDICAL ADVICE.
     WEAR SUITABLE PROTECTIVE CLOTHING.
 SECTION 4. - - - - - FIRST-AID MEASURES- - -
    IF SWALLOWED, WASH OUT MOUTH WITH WATER PROVIDED PERSON IS CONSCIOUS.
     IF INHALED, REMOVE TO PRESH AIR. IF NOT BREATHING GIVE ARTIFICIAL
     RESPINATION. IF DREATHING IS DIFFICULT, GIVE OXYGEN.
     IN CASE OF CONTACT, IMMEDIATELY WASH SKIN WITH SOAD AND COPIOUS
     IN CASE OF CONTACT, IMMEDIATELY FIGURE EYES WITH COPICUS AMOUNTS OF
     AMOUNTS OF WATER.
     WATER FOR AT LEAST 15 MINUTES.
  SECTION 5. - - - FIRE FIGHTING MEASURES - - - -
   EXTINGUISHING MEDIA
     CARSON DIOXIDE, DRY CHEMICAL POWDER OR APPROPRIATE FOAM.
    WATER EPRAY.
    SPECIAL FIREFIGHTING PROCEDURES
      WEAR SELF-CONTAINED BREATHING APPARATUS AND PROTECTIVE CLOTHING TO
      PREVENT CONTACT WITH SKIN AND EYES.
    UNUSUAL FIRE AND EXPLOSIONS HAZARDS
```

EMITS TOXIC FUNES UNDER FIRE CONDITIONS.

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WEAR RESPIRATOR, CHEMICAL SAFETY GOOGLES, RUBBER BOOTS AND HEAVY
          SWEEP UP, PLACE IN A BAG AND HOLD FOR WASTE DISPOSAL.
          RUBBER GLOVES.
          AVOID RAISING DUST.
          VENTILATE AREA AND WASH SPILL SITE AFTER MATERIAL PICKUP IS COMPLETE.
       SECTION 7. -- -- HANDLING AND STORAGE----
       SECTION 8 ---- EXPOSURE CONTROLS/PERSONAL PROTECTION----
           REFER TO SECTION 8.
        SAFETY SHOWER AND EYE BATH.
           MECHANICAL EXHAUST REQUIRED.
           WASH THOROUGHLY AFTER HANDLING.
          DO NOT BREATHE DUST.
           AVOID CONTACT WITH EYES, SKIN AND CLOTHING.
          AVOID PROLONGED OR REPEATED EXPOSURE.
         NIOSH/MSHA-APPROVED RESPIRATOR.
         COMPATIBLE CHEMICAL-RESISTANT GLOVES.
          CHEMICAL SALETY GOGGLES.
           KERP TIGHTLY CLOSED.
          STORE IN A COOL DRY PLACE.
SECTION 9. - - - - - PHYSICAL AND CHEMICAL PROPERTIES - -
          DATA NUT AVAILABLE
                    SECTION 10.
        SLYBIFTLA
           STABLE.
        incompatin (Lities
          STRONG OXIDIZING AGENTS
        HAZARDONS COMBUSTION OR DECOMPOSITION PRODUCTS
          CARBON MONOXIDE, CARBON DIOXIDE
          SULFUR OXIDES
           SODIUM OXIDES
        HAZARDOUS POLYMERIZATION
          WILL NOT OCCUR.
                           - - - TOXICOLOGICAL INFORMATION - - -
       SECTION 11. - - - - -
        ACUTE EFFECTS
         CAUSES SKIN IRRITATION.
            MAY BE HARMFUL IF ABSORBED THROUGH THE EKIN.
           CAUSES HYE IRRITATION.
           MAY BE HARMFUL IF INHALED.
           MATERIAL IS IRRITATING TO MUCOUS MEMBRANES AND UPPER
           RESPIRATORY TRACT.
          MAY BE HARMFUL IF SWALLOWED.
          TO THE HEST OF OUR KNOWLEDGE, THE CHEMICAL, PHYSICAL, AND
            TOXICOLOGICAL PROPERTIES HAVE NOT BEEN THOROUGHLY INVESTIGATED.
          RTECS #: CB1095300
           2-ANTHRACENESULFONIC ACID, 9.10-DIHYDRO-3.4-DJHYDROXY-9.10 DIOXO-.
            MCINOSODIUM SALT
          TOXICITY DATA
                                                     EXPRAM 28,780,1972
            JVN-MUS LD50:70 MG/KG
            ONLY SELECTED REGISTRY OF TOXIC EFFECTS OF CHEMICAL SUBSTANCES
            (MTECS) DATA IS PRESENTED HERE. SEE ACTUAL ENTRY IN RIECS FOR
            COMPLETE INFORMATION.
         SECTION 12. - - - - - - - ECOLOGICAL INFORMATION - - - - -
           DATA NOT YET AVAILABLE.
         SECTION 13. - . - - - - DISPOSAL CONSIDERATIONS - - - - -
            COMPACT A LICENSED PROPESSIONAL WANTE DISPOSAL SERVICE TO DISPOSE OF
            DISSOLVE OR MIX THE MATERIAL WITH A COMBUSTIBLE SOLVENT AND BURN IN A
            THIS MATERIAL.
            CHEMICAL INCINERATOR EQUIPPED WITH AN AFTERBURNER AND SCRUBBER.
            OBSERVE ALL PEDERAL, STATE AND LOCAL ENVIRONMENTAL REGULATIONS.
```

Safety (MSDS) data for bromothymol blue

General

Synonyms: 3',3"-dibromothymolsolfonphthalein

Use: indicator

Molecular formula: C27H28Br2O5S

CAS No: 76-59-5 EINECS No:

Physical data

Appearance: purple to pink powder

Melting point: 200 C

Boiling point:

Vapour density:

Vapour pressure:

Density (g cm⁻⁵):

Flash point:

Explosion limits:

Autoignition temperature:

Water solubility: negligible

Stability

Stable. Incompatible with strong oxidizing agents.

Toxicology

May act as an irritant.

Toxicity data

(The meaning of any abbreviations which appear in this section is given here.)

Risk phrases

(The meaning of any risk phrases which appear in this section is given here.)

Transport information

(The meaning of any UN hazard codes which appear in this section is given here.)

USSN 09/758,978 Torok et af. Exhibit - 11

GFS CHEMICALS, INC.

P.O. Box 245 Pewell, OH 43065 740-581-5901(ToL) 740-881-5989(Fax) 1-100-424-9300(Chuiutres 24Hr. Info.)

MATERIAL SAFETY DATA SHEET

674

SARA TITLE 313

Νo

CAS#

3051-11-4

RRILLIANT YELLOW

CHEMICAL NAME & SYNONYMS

Brilliant Yellow

FORMULA CasH11N.O1S2Naz

PHYSICAL DATA

Water soluble pH indicator range 6.0 - 8.0 yellow-orange.

<u>appearance & odor</u>

Yellow-orange powder. Odorless.

REACTIVITY & CONDITIONS TO AVOID

No hazardous self-reactivity.

FIRE HAZARDS

Will be consumed in general fire. Avoid inhalation of smoke or fumes.

EXTINGUISHER

Fight surrounding fire

DOT CLASS

NR

624.57

N/A

UEL N/A

HEALTH HAZARDS

Will stain tissue. Internal effects unknown. LD/TD no data. PHL/TLV not established. No evidence of carcinogenicity.

SPECIAL PRECAUTIONS

Use good laboratory practices. Wear goggles while handling.

Wash up thoroughly with water after contact. If swallowed, give water or milk and induce vomiting. Call a physician.

SPILLS & LEAKS

Wash up with water. Disposal to incincrator or sanitary drain.

CATALOG#

PREPARED BY

LM

DATE 7/1/96

> Exhibit - 12 USSN 09/758,978 Torok et at.

SECTION I - Product Identification	
PRODUCT NAME: BRILLIANT YELLOW FORM 624,55 CAS NO.: COMMON SYNONYMS: DIREC	ULA C26H18N4O8S2NA2 FORMULA WT:
Precautionary Labeling	CRS 3051-11-
N/A	
SECTION II - Hazardous Components	
N/A	
SECTION III - Physical Data	
BOILING POINT: N/A VAPOR PRESSURE @ 20 DENSITY (AIR=1): N/A SPECIFIC GRAVITY: N (BUTYL ACETATE=1) SOLUBILITY(H2O): SOI N/A APPEARANCE & ODOR: ORANGE POWD	OBLE I ENCOME
SECTION IV - Fire and Explosion Hazard Data	
FLASH POINT: NONE FLAMMABLE LIMITS: I EXTINGUISHING MEDIA WATER FOG, CO2, I PROCEDURES WEAR SELF-CONTAINED BRE EXPLOSION HAZARDS EMITS TOXIC FUMES	ATHING APPARATUS UNUSUAL FIRE AND
SECTION V - Health Hazard Data	
THRESHOLD LIMIT VALUE (TLV/TWA): 5 MG EFFECTS OF OVEREXPUSURE CONTACT WI IRRITATION, INHALATION OF DUST MAY R EMERGENCY AND FIRST AID PROCEDURES FOR AT LEAST 15 MINUTES SKIN: WASH AF AND WATER DUST INHALATION: REMOVE RESPIRATION IF NECESSARY INGESTION: 1	FYES: FLUSH THOROUGHT.Y WITH WATER FECTED AREAS THOROUGHLY WITH SOAP TO FRESH AIR: GIVE ARTIFICIAL
SECTION VI - Reactivity Data	
STABILITY: STABLE CONDITIONS TO AVOI POLYMERIZABLE MATERIALS DECOMPOS	D: NONE INCOMPATIBILES: OXIDIZERS, ITION PRODUCTS: COX, SOX, NOX, NH3
SECTION VII - Spill and Disposal Procedures	
STEPS TO BE TAKEN IN THE EVENT OF A S	PILL OR DISCHARGE TAKE UP &

PAGE 25/31 * RCVD AT 10/3/2005 2:55:52 PM [Eastern Daylight Time] * SVR:USPTO-EFXRF-6/25 * DNIS:2738300 * CSID:2482581439 * DURATION (mm-ss):07-14

SECTION VIII - Protective Equipment

MSDS Number: P2070 * * * * * * Effective Dute: 05/08/03 * * * * * * Supercedes: 07/13/00

25 Hour Enverowry Telephone: 069-809-9151
GHENTEID: 1-406-624-850

MSDS Material Safety Data Shoet

From: Mailinckrodt Batue, Inc.
222 Red School Lane
Philippeburg, NJ Datas

Mallinckrodt
CHEMICALS

T, Bokker

Philippeburg, NJ Datas

Mallinckrodt
II, Bokker

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All non-emergency questions should be directed to Cuctomor Sprace (1-800-862-2557) for excitatores.

PHENOL RED, SODIUM SALT

1. Product Identification

Synonyms: Phenol red, indicator grade (water soluble); phenolsulforphthalein sodium salt;

4,4'-(3H-2,1 benzoxathiol-3-ylidene) his-phonol, S,S-dioxide, sodium salt

CAS No.: 34487-61-1 Molecular Weight: 376.36

Chemical Formula: C19H14O5S Na

Product Codes: J.T. Baker: T265 Mallinckredt: 4266

2. Composition/Information on Ingredients

Ingredient	 CAS No	Percent	Hazardous
Phenol Red, Sodium Salt	34487-61-1	90 - 100%	برع برع

Exhibit - 13 USSN 09/758,978 Torok et al.

3. Hazards Identification

Emergency Overview

PAGE 26/31 * RCVD AT 10/3/2005 2:55:52 PM [Eastern Daylight Time] * SVR:USPTO-EFXRF-6/25 * DNIS:2738300 * CSID:2482581439 * DURATION (mm-ss):07-14

CAUTION! MAY DE HARMFUL IF SWALLOWED OR INHALED. MAY CAUSE IRRITATION TO SKIN, EYES, AND RESPIRATORY TRACT.

J.T. Baker SAF-T-DATA (tm) Ratings (Provided here for your convenience)

Health Rating: 1 - Slight Flammability Rating: 0 - None Reactivity Rating: 0 - None Contact Rating: 1 - Slight

Lah Protective Equip: GOGGLES; LAB COAT Storage Color Code: Orange (General Storage)

Potential Health Effects

May cause irritation to the respiratory tract. Symptoms may include coughing and shortness of breath.

Ingestion:

Large oral doses may cause irritation to the gastrointestinal tract. Ingestion effects have not been studied completely but may exhibit symptoms similar to phenolphthalein such as fall of blood pressure or an itching skin rash. May be a strong laxative.

Skin Contact:

May cause irritation with redness and pain.

Eye Contact:

May cause irritation, reduces and pain.

Chronic Exposure: No information found.

Aggravation of Pre-existing Conditions:

No information found.

4. First Aid Measures

Inhalation:

Remove to fresh air. Get medical attention for any breathing difficulty.

Induce vomiting immediately as directed by medical personnel. Never give anything by mouth to an unconscious person. Get medical attention.

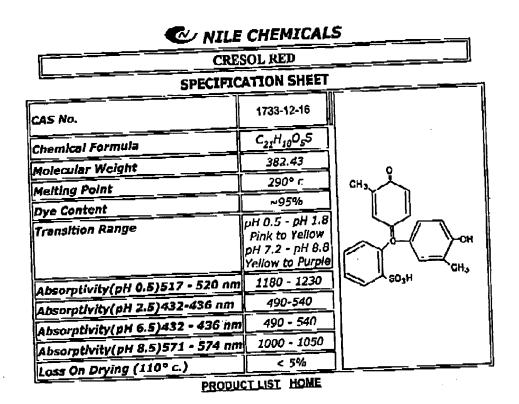
Skin Contact:

Immediately flush skin with plenty of water for at least 15 minutes. Remove contaminated clothing and shoes. Wash clothing before reuse. Thoroughly clean shoes before reuse. Get medical attention if irritation develops.

Eye Contact:

Immediately flush eyes with plenty of water for at least 15 minutes, lifting upper and lower cyclids occasionally. Get medical attention if irritation persists.

CRESOLRED



BRIEF MSDS

Hazards identification

Hazard description:- Not applicable
 Information pertaining to particular dangers for man and environment:-Not applicable

Toxicological information

Acute toxicity:
 Primary irritant effect:
 on the skin: May cause irritation
 on the eye: May cause irritation
 sensitization: No sensitizing effects known.

Transport information

Not a hazardous material for transportation.

DOT regulations:

Hazard class:
Land transport ADR/RID (cross-border)

Exhibit - 14 USSN 09/758,97 Torok et af n-Cresol Purple

MSDS Number: C5522 * * * * * Effective Date: 01/03/03 * * * * * Supercedes: 01/31/00



Material Safety Data Sheet

Fron: Mellindkrodt Baker, Inc. 222 Red School Lane Philippburg, NJ 08865



24 Hour Emergency Telephone: 908-859-2154 CHEMTREC: 1-800-424-6300

National Response in Canada CANUTEC: 613-896-8866

Outside U.S. And Canada Chembec: 703-527-3887

NOTE: CHEMTREC, CANITEC and National Response Center emergency numbers to be used only in the event of chemical emergencies involving a spill, leak, fire, exposure of accident involving chemicals.

All non-emergency questions should be directed to Customer Service (1-800-582-2537) for assistance.

m-Cresol Purple

1. Product Identification

Synonyms: m-Cresolsulfonphthalein

CAS No.: 2303-01-7

Molecular Weight: 382,44

Chemical Formula: C21H18O5S

Product Codes: F860

2. Composition/Information on Ingredients

Ingredient	CAS No	Percent	Hazardous
m-Cresol Purple	2303-01-7	90 - 100%	Yes

3. Hazards Identification

Emergency Overview

CAUTION! MAY CAUSE IRRITATION TO SKIN, EYES, AND RESPIRATORY TRACT.

Exhibit - 15 ussn 09/758,978 Torok et al.

SAF-T-DATA(tm) Ratings (Provided here for your convenience)

Health Rating: 1 - Slight Flammability Rating: 0 - None

Reactivity Rating: 0 - None Contact Rating: 1 - Slight

Lab Protective Equip: GOGGLES; LAB COAT; PROPER GLOVES

Storage Color Code: Green (General Storage)

Potential Health Effects

Inhalation:

May cause irritation to the respiratory tract.

Ingestion:

Large oral doses may cause irritation to the gastrointestinal tract.

Skin Contact:

May cause irritation with redness and pain.

Eve Contact:

May cause irritation, redness and pain.

Chronic Exposure: No information found.

Aggravation of Pre-existing Conditions:

No information found.

4. First Aid Measures

Inhalation:

Remove to fresh air. Get medical attention for any breathing difficulty.

Induce vomiting immediately as directed by medical personnel. Never give anything by mouth to an unconscious person. Get medical attention.

Immediately flush skin with plenty of soap and water for at least 15 minutes. Remove contaminated clothing and shoes. Wash clothing before reuse. Thoroughly clean shoes before reuse. Get medical attention if irritation develops.

Eye Contact:

Immediately flush eyes with plenty of water for at least 15 minutes, lifting upper and lower eyelids occasionally. Get medical attention if irritation persists.

5. Fire Fighting Measures

Not considered to be a fire hazard.

Material Safety Data Sheet Thymol Blue

ACC# 60620

Section 1 - Chemical Product and Company Identification

MSDS Name: Thymol Blue

Catalog Numbers: T416 5, T4165 Synonyms: Thymolsulfonephthalein

Company Identification: Fisher Scientific

1 Reagent Lane Fair Lawn, NJ 07410

For Information, call: 201-796-7100 Emergency Number: 201-796-7100

For CHEMTREC assistance, call: 800-424-9300

For International CHEMTREC assistance, call: 703-527-3887

Section 2 - Composition, Information on Ingredients

1	CAS#	Chemical Name	Percent	EINECS/ELINCS
	76-61-9	Thymol blue	100	<i>200-973-3</i>

Hazard Symbols: None listed. Risk Phrases: None listed.

Section 3 - Hazards Identification

EMERGENCY OVERVIEW

Appearance: green-red to brown. Caution! The toxicological properties of this material have not been fully investigated. May cause irritation.

Target Organs: None.

Potential Health Effects

Eye: May cause eye irritation. Skin: May cause skin irritation.

Ingestion: May cause irritation of the digestive tract. The toxicological properties of this substance

have not been fully investigated.

Inhalation: May cause respiratory tract irritation. The toxicological properties of this substance have not been fully investigated.

Chronic: No information found.

Section 4 - First Aid Measures

BUTZEL LONG

Suite 200 100 Bloomfield Hills Parkway Bloomfield Hills, MI 48304-2949 (248) 258-1616 R

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